

PCTWORLD INTELLECTUAL PROPERTY ORGANIZATION
International Bureau

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁷ : A61L 9 /012, 9 /14 // 101:36	A2	(11) International Publication Number: WO 00/01429 (43) International Publication Date: 13 January 2000 (13.01.00)
(21) International Application Number: PCT/GB99/01976 (22) International Filing Date: 23 June 1999 (23.06.99) (30) Priority Data: 9814372.0 2 July 1998 (02.07.98) GB (71) Applicants (for all designated States except US): UNIVERSITY OF SOUTHAMPTON [GB/GB]; Highfield, Southampton SO17 1BJ (GB). RECKITT & COLMAN PRODUCTS LIMITED [GB/GB]; One Burlington Lane, London W4 2RW (GB). (72) Inventors; and (75) Inventors/Applicants (for US only): HUGHES, John, Farrell [GB/GB]; 2 Shepherd's Close, Bartley, Southampton SO40 2JL (GB). FOX, Rodney, Thomas [GB/GB]; 30 South Street, Cottingham, Hull HU16 4AS (GB). HARRISON, Mark, Neale [GB/GB]; 27 Cromwell Close, Tutbury, Burton-on-Trent DE13 9HZ (GB). WHITMORE, Lindsey, Faye [GB/GB]; 60 Tees Farm Road, Colden Common, Winchester SO21 1UQ (GB). HARPER, Duncan, Roger [GB/GB]; Flat 2, 78 Boulevard, Hull HU3 2TA (GB). (74) Agent: ALLARD, Susan, Joyce; Boulton Wade Tennant, 27 Furnival Street, London EC4A 1PQ (GB).		(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>Without international search report and to be republished upon receipt of that report.</i>
(54) Title: TREATMENT OF AIRBORNE ALLERGENS		
(57) Abstract A method of denaturing or deactivating an airborne allergen comprising directing at the airborne source of the allergen liquid droplets from a spray device containing a liquid composition which includes an allergen denaturant or allergen deactivant, the method comprising imparting a unipolar charge to the said liquid droplets by double layer charging during the spraying of the liquid droplets by the spray device, the unipolar charge being at a level such that the said droplets have a charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg.		

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav Republic of Macedonia	TM	Turkmenistan
BF	Burkina Faso	GR	Greece			TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's Republic of Korea	NZ	New Zealand		
CM	Cameroon			PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

TREATMENT OF AIRBORNE ALLERGENS

The present invention relates to the treatment of airborne allergens.

5 Various allergens are known which are transported through the air to trigger a human reaction. For example, it has been known for a long time that house dust can trigger allergenic reactions in humans, such as asthma and rhinitis. It was reported, as early as
10 1928 that it was the dust mites in the dust that were the primary source of the allergenic response, but it was only in the 1960's that researchers appreciated its significance.

 It is believed that the faeces of the house dust
15 mite, *Dermatophogoides farinae* (known as Der-f) and *Dermatophagoides pteronyssinus* (known as Der-p) trigger the immune response of the body, thereby giving rise to well known allergenic symptoms.

 One way to overcome these allergenic responses
20 has been to vacuum clean surfaces, such as carpets, that contain the dust mites and their faeces thoroughly and often, but that is both time consuming (it has to be regularly done to ensure an allergenic free environment) and is very dependant on the efficiency
25 of the vacuum cleaner and filter bag used, e.g. micron filter bags or two layer vacuum bags.

 An alternative method of creating an allergen-free environment has been to denature the allergen, for example, by using an allergen denaturant applied
30 to airborne allergens by means of an aerosol spray device. Such a device produces an aerosol spray when

- 2 -

activated and this spray may be targeted at any space which is to be treated.

The allergens to be treated are airborne particles and the use of a known aerosol spray device results in a low collision rate between the allergen denaturant and the airborne allergens. The practical consequence of such a low collision rate is that the allergen denaturant must be used in a high amount in order to be effective. There may be other consequences such as, in the case where the aerosol spray composition includes a perfume or fragrance, a strong perfume smell or a limited fragrance choice.

Other allergens which are problematic are cat allergens (Fel-d) and cockroach allergens (Bla-g). These can be denatured using an allergen denaturant for the specific allergen applied using an aerosol spray device.

An aerosol spray type device would be of improved efficiency if the spray droplets had a greater collision rate with the allergen particles and if the droplets could wet the surface of the allergen particles. We have now developed an improved method of denaturing or deactivating airborne allergens.

According to the present there is provided a method of denaturing or deactivating an airborne allergen comprising directing at the airborne source of the allergen liquid droplets from a spray device containing a liquid composition which includes an allergen denaturant or allergen deactivant, the method comprising imparting a unipolar charge to the said liquid droplets by double layer charging during the

spraying of the liquid droplets by the spray device, the unipolar charge being at a level such that the said droplets have a charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg.

5 It is preferred that the unipolar charge which is imparted to the liquid droplets is generated solely by the interaction between the liquid within the spray device and the spray device itself as the liquid is sprayed therefrom. In particular, it is preferred
10 that the manner in which a unipolar charge is imparted to the liquid droplets does not rely even partly upon the connection of the spray device to any external charge inducing device, such as a source of relatively high voltage, or any internal charge inducing device,
15 such as a battery. With such an arrangement, the aerosol spray device is entirely self-contained making it suitable for use both in industrial, institutional and domestic situations.

 Preferably, the spray device is a domestic
20 pressure-spraying device devoid of any electrical circuitry but which is capable of being hand held. Typically such a device has a capacity in the range of from 10ml to 2000ml and can be actuated by hand, or by an automatic actuating mechanism. A particularly
25 preferred domestic device is a hand-held aerosol can.

 Preferably, therefore the droplet charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg is imparted to the liquid droplets as a result of the use of an aerosol spray device with at least one of the features of the
30 material of the actuator, the size and shape of the orifice of the actuator, the diameter of the dip tube,

the characteristics of the valve and the formulation of the allergen denaturing or allergen deactivating composition contained within the aerosol device being chosen in order to achieve the said droplet charge to mass ratio by double layer charging imparting the unipolar charge to the droplets during the actual spraying of the liquid droplets from the orifice of the aerosol spray device.

As a result of the method of the present invention there is an active targeting of allergen particles by means of the denaturant or deactivant forming part of the aerosol spray. As a result there is a perceived and actual reduction in allergenic responses due to the increase of the precipitation and deactivation of the allergen from its airborne active condition.

This result is achieved because of the unipolar charge imparted to the liquid droplets of the aerosol spray. This charge has two effects. The individual droplets are attracted to the allergen particles and, since all of the droplets carry the same polarity charge, they are repelled one from other.

Accordingly, there is little or no coalescence of the droplets and, rather, they tend to spread out to a great extent as compared to uncharged droplets. In addition, if the repulsive forces from the charge within the droplets is greater than the surface tension force of the droplets, the charged droplets are caused to fragment into a plurality of smaller charged droplets (exceeding the Rayleigh limited). This process continues until either the two opposing

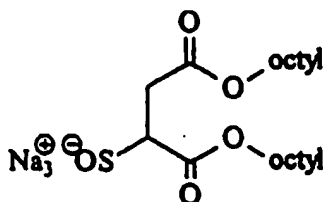
forces are equalised or the droplet has fully evaporated.

Allergen particles are normally electrically isolated from their surroundings and will typically be at a potential which is the same as that of their surroundings. An isolated allergen particle within a cloud of electrically charged liquid droplets thus is likely to cause a distortion in the configuration of the electrical field generated by the droplets so that the attraction of the droplets onto the allergen particle will be enhanced. In effect, the allergen particles are targeted by the liquid droplets.

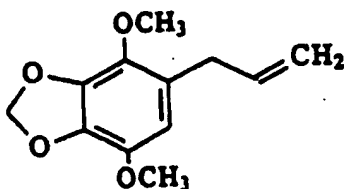
An example of an allergen denaturant is tannic acid, the use of which is described in US Patent No. 4,806,526.

Many allergen deactivants are specific to the type of dust mite allergen being treated. For example an effective Der-f allergen deactivant may not work effectively as a Der-p allergen deactivant. Various deactivants for treating Der-f and/or Der-p allergens are described in WO 99/15208.

Examples of deactivants for Der-f and/or Der-p allergens are cedarwood oil, hexadecyltrimethylammonium chloride, aluminium chlorohydrate, 1-propoxypropanol-2, polyquaternium-10, silica gel, propylene glycol alginate, ammonium sulphate, hinokitiol, L-ascorbic acid, immobilised tannic acid, chlorohexidine, maleic anhydride, hinoki oil, a composite of AgCl and TiO₂, diazolidinyl urea, 6-isopropyl-m-cresol, a compound of formula I

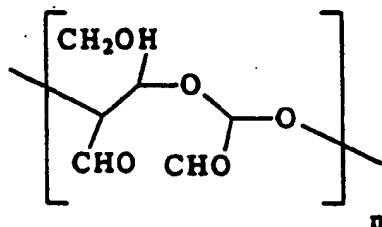


5 a compound of formula II



10

a polymeric dialdehyde containing two or more of a
recurring unit of the formula III



15

20

where $n = 2$ to 200,

urea, cyclodextrin, hydrogenated hop oil,
polyvinylpyrrolidone, N-methylpyrrolidone, the sodium
salt of anthraquinone, potassium thioglycolate or
25 glutaraldehyde.

The liquid composition which is sprayed into the
air using the aerosol spray device is preferably a
water and hydrocarbon mixture, or emulsion, or a
liquid which is converted into an emulsion by shaking
30 the spraying device before use, or during the spraying
process. An example of a composition which could be

- 7 -

prepared in a form suitable for spraying in accordance with the method of the invention is a composition based on US 4806526.

5 Whilst all liquid aerosols are known to carry a net negative or positive charge as a result of double layer charging, or the fragmentation of liquid droplets, the charge imparted to droplets of liquid sprayed from standard devices is only of the order of $\pm 1 \times 10^{-8}$ to 1×10^{-5} C/kg.

10 This invention relies on combining various characteristics of the design of an aerosol spray system so as to increase the charging of the liquid as it is sprayed from the aerosol spray device.

A typical aerosol spray device comprises:

- 15 1. An aerosol can containing the composition to be sprayed from the device and a liquid or gaseous propellant;
2. A dip tube extending into the can, the upper end of the dip tube being connected to a
- 20 valve;
3. An actuator situated above the valve which is capable of being depressed in order to operate the valve; and
4. An insert provided in the actuator
- 25 comprising an orifice from which the composition is sprayed.

A preferred aerosol spray device is described in WO 99/012227.

30 It is possible to impart higher charges to the liquid droplets by choosing aspects of the aerosol device including the material, shape and dimensions of

the actuator, the actuator insert, the valve and the dip tube and the characteristics of the liquid which is to be sprayed, so that the required level of charge is generated as the liquid is dispersed as droplets.

5 A number of characteristics of the aerosol system increase double layer charging and charge exchange between the liquid formulation and the surfaces of the aerosol system. Such increases are brought about by factors which may increase the turbulence of the flow
10 through the system, and increase the frequency and velocity of contact between the liquid and the internal surfaces of the container and valve and actuator system.

By way of example, characteristics of the
15 actuator can be optimised to increase the charge levels on the liquid sprayed from the container. A smaller orifice in the actuator insert, of a size of 0.45mm or less, increases the charge levels of the liquid sprayed through the actuator. The choice of
20 material for the actuator can also increase the charge levels on the liquid sprayed from the device with material such as nylon, polyester, acetal, PVC and polypropylene tending to increase the charge levels. The geometry of the orifice in the insert can be
25 optimised to increase the charge levels on the liquid as it is sprayed through the actuator. Inserts which promote the mechanical break-up of the liquid give better charging.

The actuator insert of the spray device may be
30 formed from a conducting, insulating, semi-conducting or static-dissipative material.

The characteristics of the dip tube can be optimised to increase charge levels in the liquid sprayed from the container. A narrow dip tube, of for example about 1.27mm internal diameter, increases the charge levels on the liquid, and the dip tube material can also be changed to increase charge.

Valve characteristics can be selected which increase the charge to mass ratio of the liquid product as it is sprayed from the container. A small tailpiece orifice in the housing, of about 0.65mm, increase product charge to mass ratio during spraying. A reduced number of holes in the stem, for example 2 x 0.50mm, also increases product charge during spray. The presence of a vapour phase tap helps to maximise the charge levels, a larger orifice vapour phase tap of, for example, about 0.50mm to 1.0mm generally giving higher charge levels.

Changes in the product formulation can also affect charging levels. A formulation containing a mixture of hydrocarbon and water, or an emulsion of an immiscible hydrocarbon and water, will carry a higher charge to mass ratio when sprayed from the aerosol device than either a water alone or hydrocarbon alone formulation.

It is preferred that an allergen neutralising composition of use in the present invention comprises an oil phase, an aqueous phase, a surfactant an allergen denaturant or allergen deactivant and a propellant.

Preferably the oil phase includes a $C_9 - C_{12}$ hydrocarbon which is preferably present in the

composition in the amount of from 2 to 10% w/w.

Preferably the surfactant is glyceryl oleate or a polyglycerol oleate, preferably present in the composition in an amount of from 0.1 to 1.0% w/w.

5 Preferably the propellant is liquified petroleum gas (LPG) which is preferably butane, optionally in admixture with propane. The propellant may be present in an amount of from 10 to 90% w/w depending upon whether the composition is intended for spraying as a
10 "wet" or as a "dry" composition. For a "wet" composition, the propellant is preferably present in an amount of from 20 to 50% w/w, more preferably in an amount of from 30 to 40% w/w.

15 The liquid droplets sprayed from the aerosol spray device will generally have diameter in the range of from 5 to 100 micrometers, with a peak of droplets of about 40 micrometers. The liquid which is sprayed from the aerosol spray device may contain a predetermined amount of a particulate material, for
20 example, fumed silica, or a predetermined amount of a volatile solid material, such as menthol or naphthalene.

25 The method of the present invention, in addition to neutralising allergens also accelerates the natural process of precipitation of airborne particles by indirect charging of the particles, thereby enabling the air quality to be improved quickly and conveniently.

30 A can for a typical spray device is formed of aluminum or lacquered or unlacquered tin plate or the like. The actuator insert may be formed of, for

instance, acetal resin. The valve stem lateral opening may typically be in the form of two apertures of diameters 0.51mm.

The present invention will now be described, by way of example only, with reference to the accompanying drawings, which:-

Figure 1 is a diagrammatic cross section through an aerosol spray apparatus in accordance with the invention;

Figure 2 is a diagrammatic cross section through the valve assembly of the apparatus of Figure 1;

Figure 3 is a cross section through the actuator insert of the assembly shown in Figure 2;

Figure 4 shows the configuration of the bore of the spraying head shown in Figure 3 when viewed in the direction A;

Figure 5 shows the configuration of the swirl chamber of the spraying head shown in Figure 3 when viewed in the direction B; and

Figure 6 illustrates the method of the invention as described in Example 6 in relation to the rate of precipitation of dust particles containing allergens.

Referring to Figures 1 and 2, an aerosol spray device in accordance with the invention is shown. It comprises a can 1, formed of aluminum or lacquered or unlacquered tin plate or the like in conventional manner, defining a reservoir 2 for a liquid 3 having a conductivity such that droplets of the liquid can carry an appropriate electrostatic charge. Also located in the can is a gas under pressure which is capable of forcing the liquid 3 out of the can 1 via a

- 12 -

conduit system comprising a dip tube 4 and a valve and actuator assembly 5. The dip tube 4 includes one end 6 which terminates at a bottom peripheral part of the can 1 and another end 7 which is connected to a tailpiece 8 of the valve assembly. The tailpiece 8 is secured by a mounting assembly 9 fitted in an opening in the top of the can and includes a lower portion 10 defining a tailpiece orifice 11 to which end 7 of the dip tube 4 is connected. The tailpiece includes a bore 12 of relatively narrow diameter at lower portion 11 and a relatively wider diameter at its upper portion 13. The valve assembly also includes a stem pipe 14 mounted within the bore 12 of the tailpiece and arranged to be axially displaced within the bore 12 against the action of spring 15. The valve stem 14 includes an internal bore 16 having one or more lateral openings (stem holes) 17 (see Figure 2). The valve assembly includes an actuator 18 having a central bore 19 which accommodates the valve stem 14 such that the bore 16 of the stem pipe 14 is in communication with bore 19 of the actuator. A passage 20 in the actuator extending perpendicularly to the bore 19 links the bore 19 with a recess including a post 21 on which is mounted a spraying head in the form of an insert 22 including a bore 23 which is in communication with the passage 20.

A ring 24 of elastomeric material is provided between the outer surface of the valve stem 14 and, ordinarily, this sealing ring closes the lateral opening 17 in the valve stem 14. The construction of the valve assembly is such that when the actuator 18

- 13 -

is manually depressed, it urges the valve stem 14 downwards against the action of the spring 15 as shown in Figure 2 so that the sealing ring 24 no longer closes the lateral opening 17. In this position, a
5 path is provided from the reservoir 2 to the bore 23 of the spraying head so that liquid can be forced, under the pressure of the gas in the can, to the spraying head via a conduit system comprising the dip tube 4, the tailpiece bore 12, the valve stem bore 16,
10 the actuator bore 19 and the passage 20.

An orifice 27 (not shown in Figure 1) is provided in the wall of the tailpiece 8 and constitutes a vapour phase tap whereby the gas pressure in the reservoir 2 can act directly on the liquid flowing
15 through the valve assembly. This increases the turbulence of the liquid. It has been found that an increased charge is provided if the diameter of the orifice 27 is at least 0.76mm.

Preferably the lateral opening 17 linking the
20 valve stem bore 16 to the tailpiece bore 12 is in the form of 2 orifices each having a diameter of not more than 0.51mm to enhance electrostatic charge generation. Further, the diameter of the dip tube 4 is preferably as small as possible, for example,
25 1.2mm, in order to increase the charge imparted to the liquid. Also, charge generation is enhanced if the diameter of the tailpiece orifice 11 is as small as possible eg not more than about 0.64mm.

Referring now to Figure 3, there is shown on an
30 increased scale, a cross section through the actuator insert of the apparatus of Figures 1 and 2. For

simplicity, the bore 23 is shown as a single cylindrical aperture in this Figure. However, the bore 23 preferably has the configuration, for instance, shown in Figure 4. The apertures of the bore 23 are denoted by reference numeral 31 and the aperture-defining portions of the bore are denoted by reference numeral 30. The total peripheral length of the aperture-defining portions at the bore outlet is denoted by L (in mm) and a is the total area of the aperture at the bore outlet (in mm^2) and the values for L and a are as indicated in Figure 4. L/a exceeds 8 and this condition has been found to be particularly conducive to charge development because it signifies an increased contact area between the actuator insert and the liquid passing there through.

Many different configurations can be adopted in order to produce a high L/a ratio without the cross-sectional area a being reduced to a value which would allow only low liquid flow rates. Thus, for example it is possible to use actuator insert bore configurations (i) wherein the bore outlet comprises a plurality of segment-like apertures (with or without a central aperture); (ii) wherein the outlet comprises a plurality of sector-like apertures; (iii) wherein the aperture together form an outlet in the form of a grill or grid; (iv) wherein the outlet is generally cruciform; (v) wherein the apertures together define an outlet in the form of concentric rings; and combinations of these configurations. Particularly preferred are actuator insert bore configurations wherein a tongue like portion protrudes into the

liquid flow stream and can be vibrated thereby. This vibrational property may cause turbulent flow and enhanced electrostatic charge separation of the double layer, allowing more charge to move into the bulk of the liquid.

Referring now to Figure 5, there is shown a plan view of one possible configuration of swirl chamber 35 of the actuator insert 22. The swirl chamber includes 4 lateral channels 36 equally spaced and tangential to a central area 37 surrounding the bore 23. In use, the liquid driven from the reservoir 2 by the gas under pressure travels along passage 20 and strikes the channels 36 normal to the longitudinal axis of the channels. The arrangement of the channels is such that the liquid tends to follow a circular motion prior to entering the central area 37 and thence the bore 23. As a consequence, the liquid is subjected to substantial turbulence which enhances the electrostatic charge in the liquid.

The following Examples illustrate the invention:-

EXAMPLE 1

The concentration of allergen (for example Der p1, Der f1, Fel d1 or Bla g1) in an artificially created aerosol of dust particles (of domestic origin) was quantified, and compared with the configuration following treatment of the dust cloud with a charged liquid aerosol containing a neutralising agent and an identical charged liquid aerosol not containing a neutralising agent. The dust cloud was generated in a

- 16 -

test chamber of 2.2m³ by dispersing 2.0g of house dust with compressed air. 2.0g of liquid aerosol (either with or without the neutralising agent) were sprayed into the centre of the dust cloud, and air sampling
5 commenced immediately. Air was sampled at a rate of 18 liters per minute for 5 minutes, through a glass fibre filter paper supported in an in-line filter holder (German laboratories) to collect airborne particles. Allergen was eluted from these collection
10 papers in 1 ml of 10% BSA PBS-T (Phosphate buffered Saline with 0.05% Tween and 10% Bovine Serum Albumen) overnight. The filter paper was then removed and the remaining solution centrifuged for 5 minutes at 13,000rpm. The supernatant, containing the allergen
15 in solution, was decanted into a clean container. Control measurements were taken by sampling the dust cloud without treating it with liquid aerosol. A minimum of 5 replicates were performed for the control and for treatment with the charged aerosol containing
20 a neutralising agent and the equivalent aerosol without the neutralising agent. Allergen concentrations in the solutions collected were assayed using standard ELISA (Enzyme Linked ImmunoSorbent Assay) methods.

25 The aerosol formulation containing a neutralising agent was produced from the following ingredients:

Ethyl Alcohol (30% v/v)

Water (59% v/v)

Benzyl Alcohol (10% v/v)

30 Tannic Acid (1% w/v)

The mixture was introduced into a conventional

aerosol can.

The can was pressurized using compressed air to achieve a pressure of 130 psi within the can.

5 The charge level on droplets emitted from this spray can was artificially raised to a charge to mass ratio of -1×10^{-4} C/kg by supplying -10 kv charge to the seam of the can from a high voltage power supply. A flow rate of approximately 1.5 g/sec was obtained. The droplets became rapidly dispersed in the air.

10 The above-described aerosol spray device was compared with a standard, known aerosol spray device loaded with the same aerosol formulation. When used to counteract allergens in a room according to the protocol as described above, it was found that the
15 amount of spraying required was significantly less with the above-described device compared with the standard, known device.

EXAMPLE 2

20

The depletion rate of dust particles in an actinically created dust cloud containing a quantified concentration of allergen was quantified using an air particle counter (APC 300A, Malvern Instruments,
25 Malvern, UK). The rate of depletion occurring through natural processes alone was compared with the rate after treatment of the dust cloud with a charged liquid aerosol composition containing an allergen denaturant (charge-to-mass ratio of -1.4×10^{-4} C/kg⁻¹)
30 sprayed from a hand-held pressure-pack dispenser, and an equivalent liquid aerosol which was not charged

(charge-to-mass ratio of $- 1.3 \times 10^{-6} \text{ C/kg}^{-1}$). A cloud of dust particles (of domestic origin) was created in a test chamber 2.2m^3 , by dispersing 2.0g of dust with compressed air. The concentration of particles in this cloud over a 14 minute period was quantified using the air particle counter, and gave a rate for the natural depletion rate for this dust. To quantify the effect of the charged and uncharged liquid aerosol sprays on the concentration of dust particles a 2.0g spray of one of these sprays was made into the center of the artificial dust cloud, immediately following a first measurement of particle concentration. Subsequent measurements of dust concentration reflected the particle depletion achieved by the liquid aerosol spray. A minimum of 5 replicates were conducted for the natural rate of particle settling, and for depletion caused by the uncharged and the charged liquid aerosol spray.

Typical results are shown in Figure 6 for the percentage of 1 to 2 micron diameter particles remaining airborne. The rate at which dust particles settle due to natural processes alone was quite steady and slow. Liquid aerosol sprays, generated by hand-held pressure-pack devices caused the depletion of 60% of the particles within 90 seconds. Charged liquid aerosol sprays depleted almost 80% of particles within 90 seconds. The improved particle depletion and allergen denaturation achieved when the liquid aerosol was charged over that achieved by the uncharged liquid aerosol was statistically significant ($p < 0.05$).

CLAIMS:

1. A method of denaturing or deactivating an airborne allergen comprising directing at the airborne source of the allergen liquid droplets from a spray device containing a liquid composition which includes an allergen denaturant or allergen deactivant, the method comprising imparting a unipolar charge to the said liquid droplets by double layer charging during the spraying of the liquid droplets by the spray device, the unipolar charge being at a level such that the said droplets have a charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg.

2. A method as claimed in claim 1 wherein the spray device is an aerosol spray device.

3. A method as claimed in claim 1 or claim 2 wherein the liquid composition is an emulsion.

4. A method as claimed in any one of the preceding claims wherein the liquid droplets have a diameter in the range of from 5 to 100 micrometers.

5. A method as claimed in any one of the preceding claims wherein the composition includes an allergen denaturant or deactivant effective against Der-f, Der-p, Fel-d and/or Bla-g allergens.

6. A method as claimed in any one of the preceding claims wherein the unipolar charge is

imparted to the liquid droplets solely by the interaction between the liquid and the spray device, without any charge being imparted thereto from an internal or external charge inducing device.

5

7. A method as claimed in claim 6 wherein the droplet charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg is imparted to the liquid droplets as a result of the use of an aerosol spray device with at least one of the features of the material of the actuator, the size and shape of the orifice of the actuator, the diameter of the dip tube, the characteristics of the valve and the formulation of the allergen denaturing or allergen deactivating composition contained within the aerosol device being chosen in order to achieve the said droplet charge to mass ratio by double layer charging imparting the unipolar charge to the droplets during the actual spraying of the liquid droplets from the orifice of the aerosol spray device.

20

8. A method as claimed in any one of the preceding claims wherein the liquid composition comprises an oil phase, an aqueous phase, a surfactant, an allergen denaturant or allergen deactivant and a propellant.

25

9. A method as claimed in claim 8 wherein the oil phase includes a $C_9 - C_{12}$ hydrocarbon.

30

10. A method as claimed in claim 9 wherein the $C_9 - C_{12}$ hydrocarbon is present in the composition in

an amount of from 2 to 10% w/w.

11. A method as claimed in any one of claims 8 to 10 wherein the surfactant is glyceryl oleate or a polyglycerol oleate.

12. A method as claimed in any one of claims 8 to 11 wherein the surfactant is present in the composition in an amount of from 0.1 to 1.0% w/w.

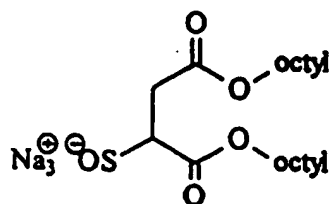
10

13. A method as claimed in any one of claims 8 to 12 wherein the propellant is liquified petroleum gas or compressed gas.

14. A method as claimed in claim 13 wherein the propellant is present in the composition in an amount of from 20 to 50% w/w.

15. A method as claimed in any one of the preceding claims wherein the allergen denaturant is tannic acid, Examples of deactivants for Der-f and/or Der-p allergens are cedarwood oil, hexadecyltrimethylammonium chloride, aluminium chlorohydrate, 1-propoxypropanol-2, polyquaternium-10, silica gel, propylene glycol alginate, ammonium sulphate, hinokitiol, L-ascorbic acid, immobilised tannic acid, chlorohexidine, maleic anhydride, hinoki oil, a composite of AgCl and TiO₂, diazolidinyl urea, 6-isopropyl-m-cresol, a compound of formula I

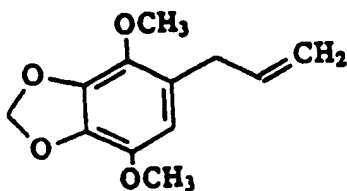
30



5

a compound of formula II

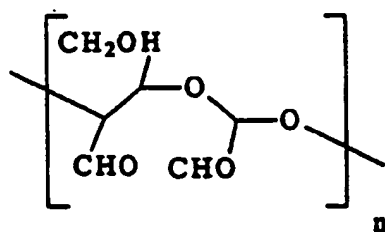
10



15

a polymeric dialdehyde containing two or more of a
recurring unit of the formula III

20

where $n = 2$ to 200,

25

urea, cyclodextrin, hydrogenated hop oil,
polyvinylpyrrolidone, N-methylpyrrolidone, the sodium
salt of anthraquinone, potassium thioglycolate or
glutaraldehyde.

FIG. 1

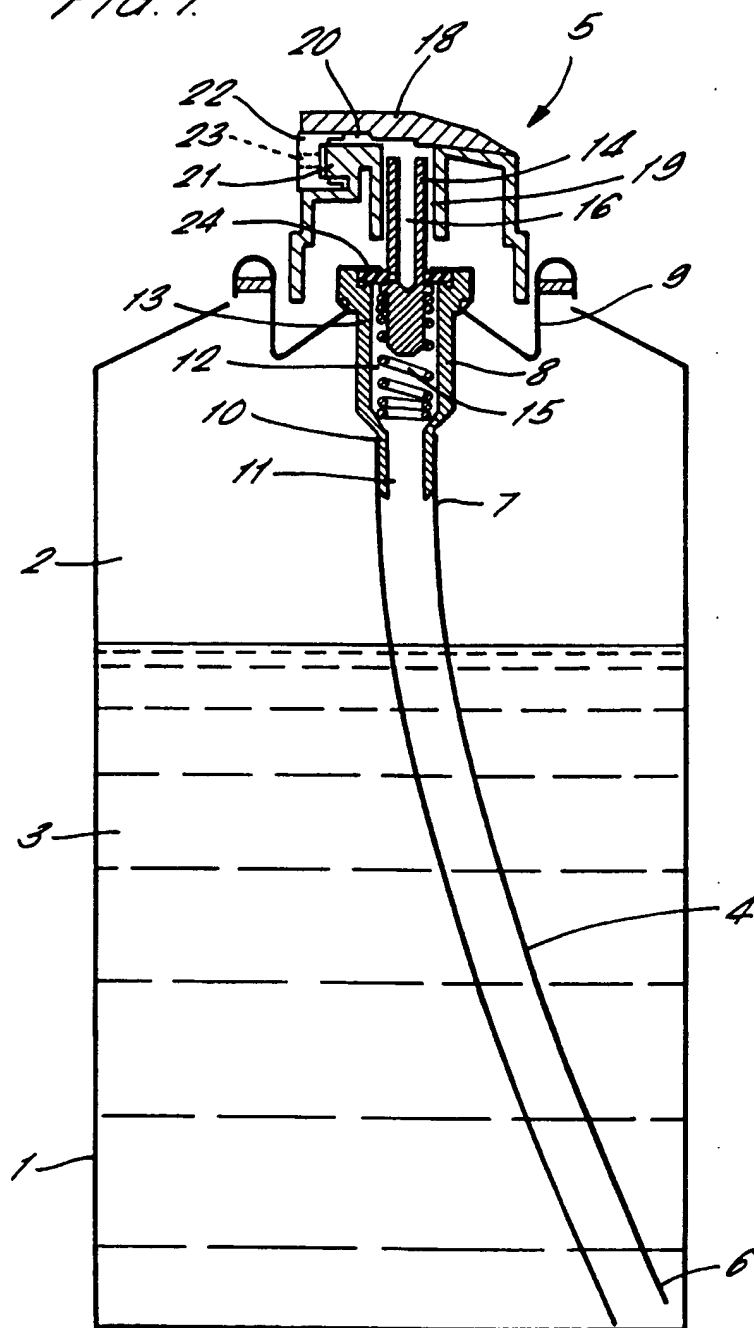
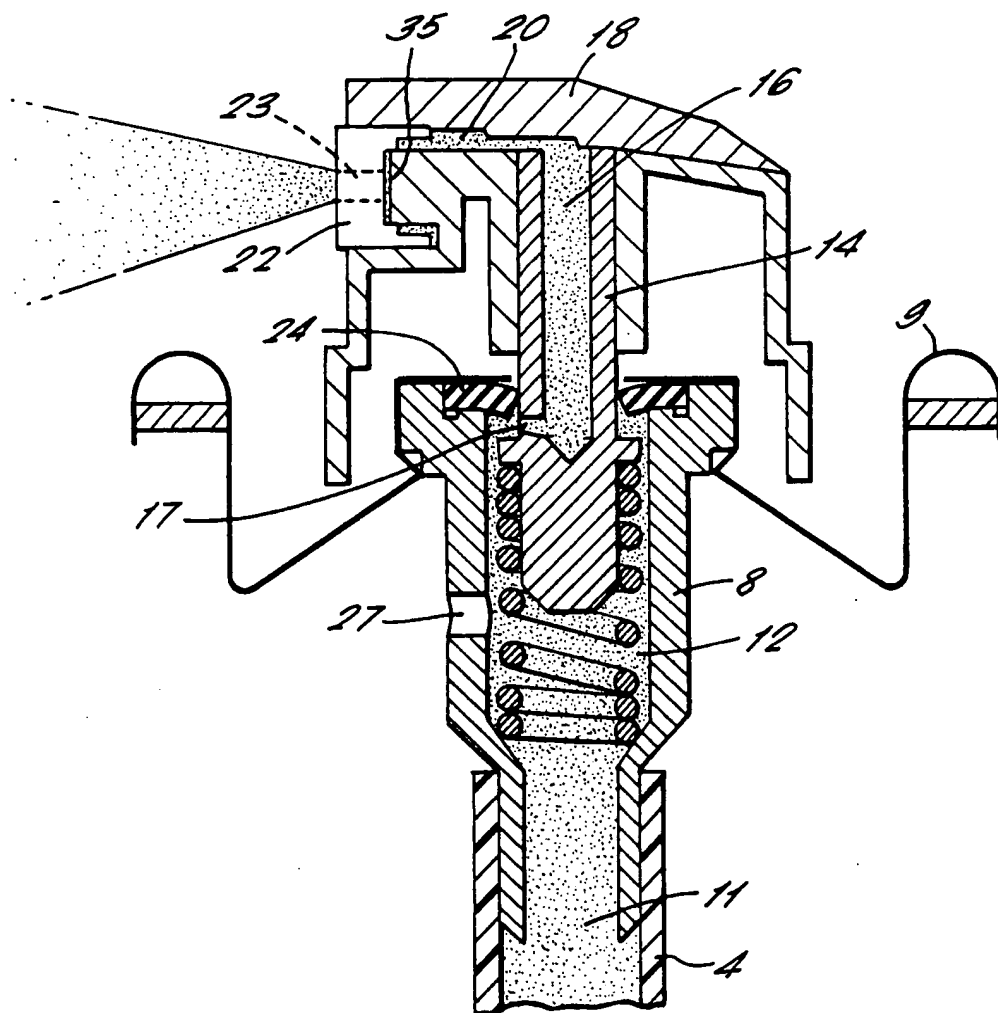


FIG. 2.



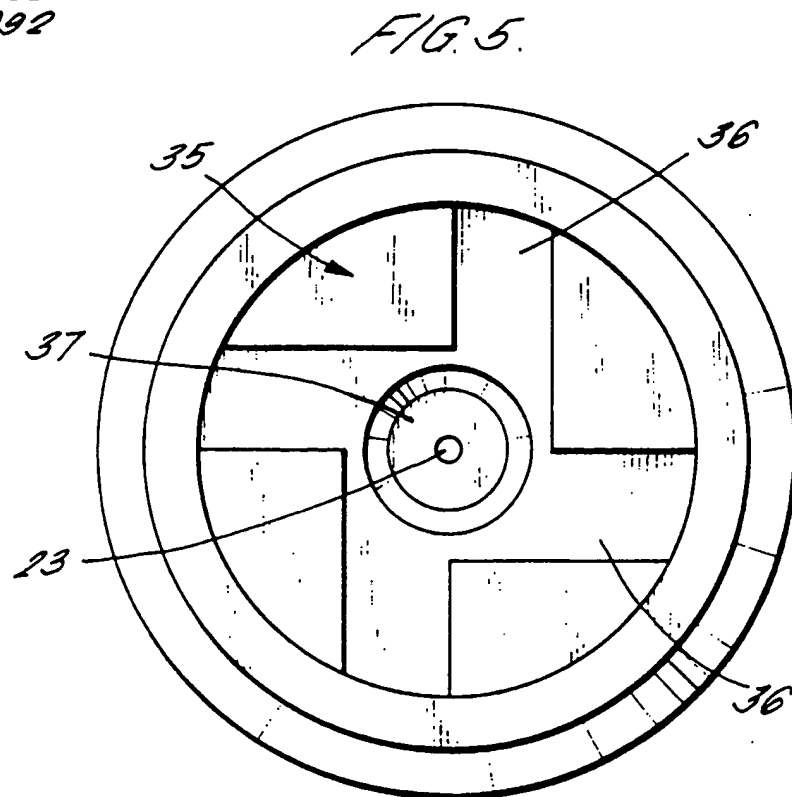
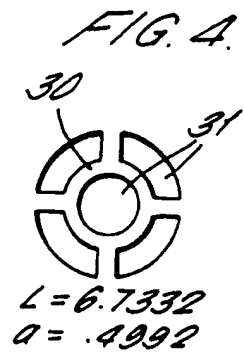
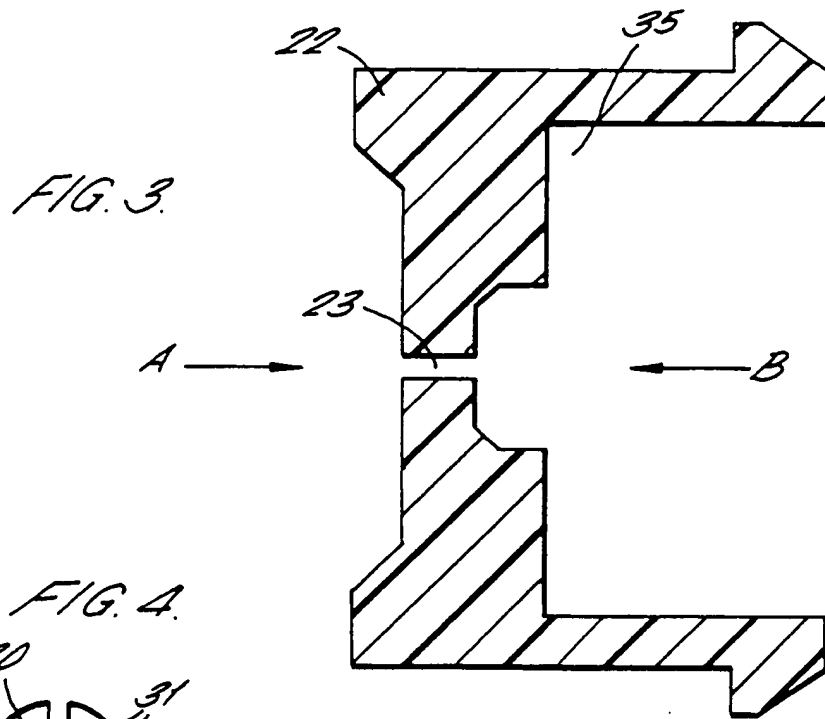
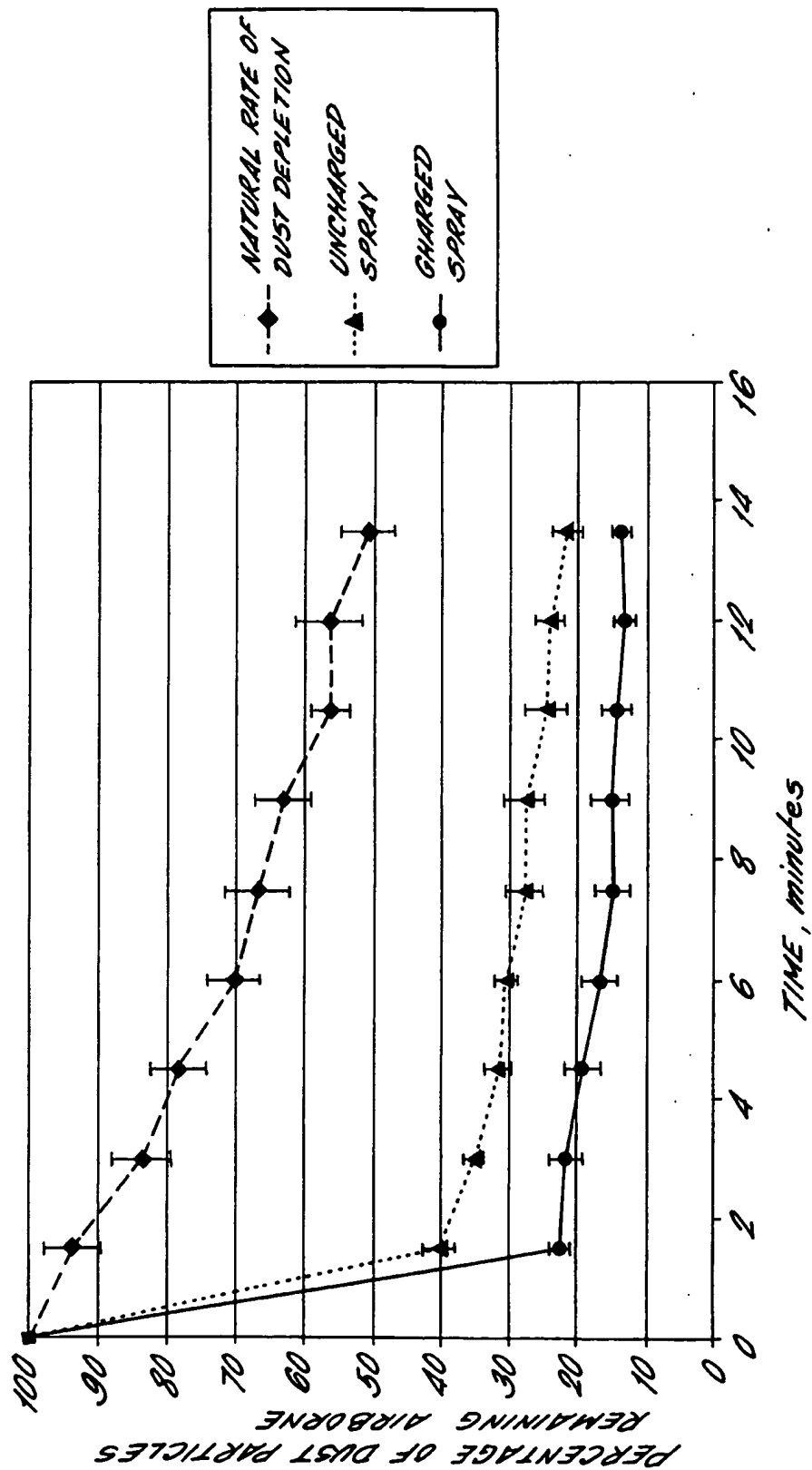


FIG. 6.





INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁷ : A61L 9/14, 9/01	A3	(11) International Publication Number: WO 00/01429 (43) International Publication Date: 13 January 2000 (13.01.00)
(21) International Application Number: PCT/GB99/01976 (22) International Filing Date: 23 June 1999 (23.06.99) (30) Priority Data: 9814372.0 2 July 1998 (02.07.98) GB (71) Applicants (for all designated States except US): UNIVERSITY OF SOUTHAMPTON [GB/GB]; Highfield, Southampton SO17 1BJ (GB). RECKITT & COLMAN PRODUCTS LIMITED [GB/GB]; One Burlington Lane, London W4 2RW (GB). (72) Inventors; and (75) Inventors/Applicants (for US only): HUGHES, John, Farrell [GB/GB]; 2 Shepherd's Close, Bartley, Southampton SO40 2JL (GB). FOX, Rodney, Thomas [GB/GB]; 30 South Street, Cottingham, Hull HU16 4AS (GB). HARRISON, Mark, Neale [GB/GB]; 27 Cromwell Close, Tutbury, Burton-on-Trent DE13 9HZ (GB). WHITMORE, Lindsey, Faye [GB/GB]; 60 Tees Farm Road, Colden Common, Winchester SO21 1UQ (GB). HARPER, Duncan, Roger [GB/GB]; Flat 2, 78 Boulevard, Hull HU3 2TA (GB). (74) Agent: ALLARD, Susan, Joyce; Boulton Wade Tennant, 27 Furnival Street, London EC4A 1PQ (GB).		(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>With international search report.</i> <i>Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</i> (88) Date of publication of the international search report: 6 April 2000 (06.04.00)
(54) Title: TREATMENT OF AIRBORNE ALLERGENS (57) Abstract A method of denaturing or deactivating an airborne allergen comprising directing at the airborne source of the allergen liquid droplets from a spray device containing a liquid composition which includes an allergen denaturant or allergen deactivant, the method comprising imparting a unipolar charge to the said liquid droplets by double layer charging during the spraying of the liquid droplets by the spray device, the unipolar charge being at a level such that the said droplets have a charge to mass ratio of at least $\pm 1 \times 10^{-4}$ C/kg.		

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav Republic of Macedonia	TM	Turkmenistan
BF	Burkina Faso	GR	Greece	ML	Mali	TR	Turkey
BG	Bulgaria	HU	Hungary	MN	Mongolia	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MR	Mauritania	UA	Ukraine
BR	Brazil	IL	Israel	MW	Malawi	UG	Uganda
BY	Belarus	IS	Iceland	MX	Mexico	US	United States of America
CA	Canada	IT	Italy	NE	Niger	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NL	Netherlands	VN	Viet Nam
CG	Congo	KE	Kenya	NO	Norway	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NZ	New Zealand	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's Republic of Korea	PL	Poland		
CM	Cameroon	KR	Republic of Korea	PT	Portugal		
CN	China	KZ	Kazakhstan	RO	Romania		
CU	Cuba	LC	Saint Lucia	RU	Russian Federation		
CZ	Czech Republic	LI	Liechtenstein	SD	Sudan		
DE	Germany	LK	Sri Lanka	SE	Sweden		
DK	Denmark	LR	Liberia	SG	Singapore		
EE	Estonia						

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 99/01976

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 A61L9/14 A61L9/01

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A61L

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	WO 98 24356 A (GAYNOR PAUL TERENCE ;HUGHES JOHN FARRELL (GB); MCKECHNIE MALCOLM T) 11 June 1998 (1998-06-11) page 2, line 29 - line 34 page 4, line 12 - line 27; claim 1 page 5, line 6 - line 13 page 9, line 30 - line 35 ---	1-4, 15
Y	WO 97 28883 A (FOX RODNEY THOMAS ;HARRISON NEALE MARK (GB); HUGHES JOHN FARRELL () 14 August 1997 (1997-08-14) page 1, line 15 - line 29 page 2, line 27 - line 35 page 4, line 1 - line 26 ---	1-4
Y	EP 0 168 243 A (UNIVERSITY OF SYDNEY) 15 January 1986 (1986-01-15) page 4, line 6; claim 1 ---	15
-/--		

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

* Special categories of cited documents :

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Date of the actual completion of the international search

7 February 2000

Date of mailing of the international search report

14/02/2000

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2
NL - 2280 HV Rijswijk
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,
Fax: (+31-70) 340-3016

Authorized officer

Peltre, C

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 99/01976

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>US 4 776 515 A (MICHALCHIK M.) 11 October 1988 (1988-10-11) column 2, line 57 - line 61 column 3, line 43 - line 58 column 4, line 3 - line 8 -----</p>	1

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/GB 99/01976

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9824356 A	11-06-1998	GB 2328862 A AU 5401798 A EP 0942680 A	10-03-1999 29-06-1998 22-09-1999
WO 9728883 A	14-08-1997	AU 1613397 A BR 9707383 A CN 1214640 A EP 0888162 A NZ 331255 A	28-08-1997 13-04-1999 21-04-1999 07-01-1999 30-08-1999
EP 168243 A	15-01-1986	AU 577874 B AU 4460685 A AT 50144 T JP 1683570 C JP 2016731 B JP 61044821 A US 4977142 A US 4806526 A	06-10-1988 16-01-1986 15-02-1990 31-07-1992 18-04-1990 04-03-1986 11-12-1990 21-02-1989
US 4776515 A	11-10-1988	NONE	